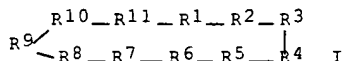


L35 ANSWER 220 OF 272 HCAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1990:77967 HCAPLUS Full-text
 DOCUMENT NUMBER: 112:77967
 TITLE: Preparation of cyclosporin analogs as
 non-immunosuppressive antagonists of cyclosporin A
 binding to cyclophilin
 INVENTOR(S): Dumont, Francis J.; Durette, Philippe L.; Pessolano,
 Arsenio A.; Boger, Joshua S.; Sigal, Nolan H.
 PATENT ASSIGNEE(S): Merck and Co., Inc., USA
 SOURCE: Brit. UK Pat. Appl., 40 pp.
 CODEN: BAXXDU
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
GB 2212499	A1	19890726	GB 1988-26033	19881107
GB 2212499	B2	19920408		
US 4914188	A	19900403	US 1987-121827	19871116
US 5236899	A	19930817	US 1991-752169	19910823
PRIORITY APPLN. INFO.:			US 1987-121827	A 19871116
			US 1989-425262	B1 19891023
OTHER SOURCE(S):		CASREACT 112:77967; MARPAT 112:77967		
GI				



AB The title compds. [I; R1 = (dihydro)MeBMT; R2 = Abu, NHCH(CO)R12; R3 = Sar, (fluorinated) D-MeAla; R4, R9, R10 = (fluorinated) MeLeu; R5 = (fluorinated) Val; R6 = (fluorinated) MeAla, (fluorinated) MeAbu; Abu = 2-aminobutyric acid residue; BMT = (4R)-4-[(E)-2-butenyl]-4-methyl-L- threonine residue; Sar = MeNCH2CO], excellent blockers of cyclosporin A binding to cyclophilin, are prepared Cyclo[(2S,3R,4R,6E)-3-hydroxy-4-methyl- 2-methylamino-6-octenoyl-L-2-aminobutyrylsarcosyl-N-methyl-L-leucyl-valyl- N-methyl-L-alanyl-D-alanyl-N-methyl-L-leucyl-N-methyl-L-leucyl-N-methyl-L-valyl] was prepared in 14 steps by the solution method starting from tert-butoxycarbonyl-N-methyl-L-alanine and L-alanine benzyl ester tosylate.

IT 111710-61-3P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of, as cyclosporin analog as non-immunosuppressive antagonist of cyclosporin A binding to cyclophilin)

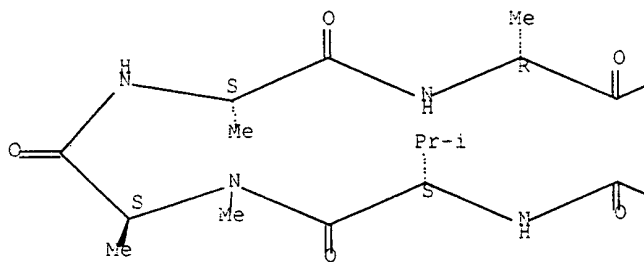
RN 111710-61-3 HCAPLUS

CN Cyclosporin A, 11-(N-methyl-L-alanine)- (9CI) (CA INDEX NAME)

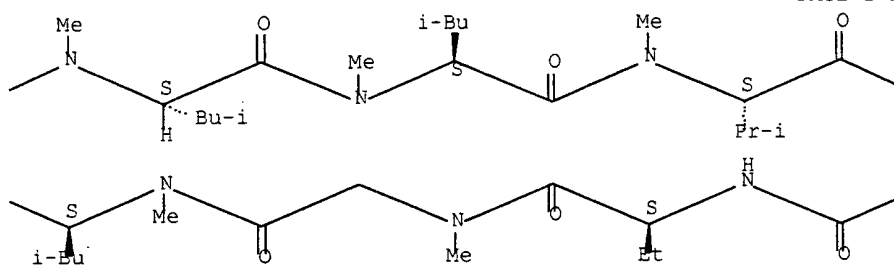
Absolute stereochemistry.

Double bond geometry as shown.

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